

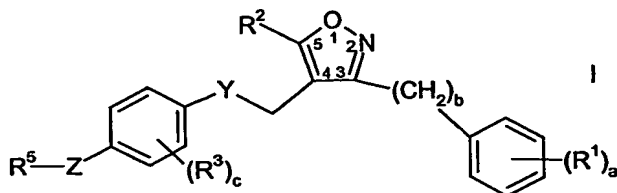
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## CLAIMS

That Which Is Claimed Is:

1. A compound of formula (I):

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wherein:

a is 1-5;

10 each  $R^1$  is the same or different and is independently selected from the group consisting of halo, alkyl, alkenyl,  $-OR^6$ ,  $-S(O)_rR^6$ ,  $-NR^6R^7$ ,  $-R^4OR^6$ ,  $-R^4S(O)_rR^6$ ,  $-R^4NR^6R^7$  and cyano;

b is 0-3;

 $R^2$  is selected from the group consisting of alkyl, alkenyl,  $C_{3-6}$ cycloalkyl,

15  $C_{3-6}$ cycloalkenyl,  $-OR^6$ ,  $-NR^6R^7$ ,  $-R^4OR^6$ ,  $-R^4NR^6R^7$ , cyano and nitro;

Y is  $-O-$  or  $-N(R^8)-$ ;

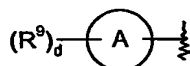
c is 0-4;

each  $R^3$  is the same or different and is independently selected from the group

20 consisting of halo, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl,  $-OR^6$ ,  $-COR^6$ ,  $-CO_2R^6$ ,  $-CH(R^6)OR^7$ ,  $-S(O)_rR^6$ ,  $-NR^6R^7$ ,  $-R^4$ cycloalkyl,  $-R^4OR^6$ ,  $-R^4COR^6$ ,  $-R^4CO_2R^6$ ,  $-R^4S(O)_rR^6$ ,  $-R^4NR^6R^7$  and cyano

Z is selected from the group consisting of  $-O-R^4-$ ,  $-R^4-O-$ ,  $-S(O)_rR^4-$ ,  $-R^4-S(O)_r-$ ,  $-N(R^8)-R^4-$ ,  $-R^4-N(R^8)-$ ,  $-C(O)N(R^8)-$ ,  $-C(O)R^4N(R^8)-$ ,  $-S(O)_rN(R^8)-$  and  $-S(O)_rR^4N(R^8)-$ ;

25 each  $R^4$  is the same or different and is independently selected from the group consisting of alkylene and alkenylene;

 $R^5$  is selected from the group consisting of  $R^6O-$ ,  $R^6O_2C-$ , and

30 wherein Ring A is aryl or a 5-12 membered heterocycle or heteroaryl;  
d is 0-4;

each R<sup>9</sup> is the same or different and is independently selected from the group consisting of halo, alkyl, alkenyl, alkynyl, cycloalkyl, -OR<sup>6</sup>, -COR<sup>6</sup>, -CO<sub>2</sub>R<sup>6</sup>, -CH(R<sup>6</sup>)OR<sup>7</sup>, -S(O)<sub>r</sub>R<sup>6</sup>, -NR<sup>6</sup>R<sup>7</sup>, -R<sup>4</sup>cycloalkyl, -R<sup>4</sup>OR<sup>6</sup>, -R<sup>4</sup>COR<sup>6</sup>, -R<sup>4</sup>CO<sub>2</sub>R<sup>6</sup>, -R<sup>4</sup>S(O)<sub>r</sub>R<sup>6</sup>, -R<sup>4</sup>NR<sup>6</sup>R<sup>7</sup>, cyano, 5-9 membered heterocycle and 5-9 membered heteroaryl;

each R<sup>6</sup> and R<sup>7</sup> are the same or different and are each independently selected from the group consisting of H, alkyl, alkenyl, C<sub>3-6</sub>cycloalkyl and C<sub>3-6</sub>cycloalkenyl; R<sup>8</sup> is H or alkyl; and

each f is the same or different and is independently selected from the group consisting of 0, 1 and 2;

or a pharmaceutically acceptable salt, solvate or physiologically functional derivative thereof.

2. The compound according to claim 1 wherein a is 1-2.

3. The compound according to any of claims 1-2 wherein each R<sup>1</sup> is the same or different and is independently selected from the group consisting of halo and -OR<sup>6</sup>.

4. The compound according to any of claims 1-3 wherein b is 0 or 1.

5. The compound according to any of claims 1-4 wherein R<sup>2</sup> is selected from the group consisting of alkyl and C<sub>3-6</sub>cycloalkyl.

6. The compound according to any of claims 1-5 wherein Y is -O-.


7. The compound according to any of claims 1-6, wherein c is 0-2.


8. The compound according to any of claims 1-7, wherein each R<sup>3</sup> is the same or different and is independently selected from the group consisting of halo and alkyl.

9. The compound according to any of claims 1-8, wherein Z is selected from the group consisting of -O-R<sup>4</sup>-, -R<sup>4</sup>-O-, -S(O)<sub>r</sub>-R<sup>4</sup>-, -N(R<sup>8</sup>)-R<sup>4</sup>-, -R<sup>4</sup>-N(R<sup>8</sup>)-, -C(O)N(R<sup>8</sup>)-,

$-\text{C}(\text{O})\text{R}^4\text{N}(\text{R}^8)-$ ,  $-\text{S}(\text{O})_r\text{N}(\text{R}^8)-$  and  $-\text{S}(\text{O})_r\text{R}^4\text{N}(\text{R}^8)-$ .

10. The compound according to any of claims 1-9, wherein  $\text{R}^8$  is H or methyl.

5 11. The compound according to any of claims 1-10, wherein  $\text{R}^5$  is selected from the group consisting of  $\text{R}^6\text{O}_2\text{C}-$ , and  $(\text{R}^9)_d-\text{A}$  .

12. The compound according to any of claims 1-11, wherein  $\text{R}^5$  is  $(\text{R}^9)_d-\text{A}$   and Ring A is phenyl or furan.

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13. A pharmaceutical composition comprising a compound according to any of claims 1-12.

14. The pharmaceutical composition according to claim 13, further comprising a  
15 pharmaceutically acceptable carrier or diluent.

15. A method for the treatment or prophylaxis of a condition mediated by FXR in a subject, said method comprising administering to said subject a therapeutically effective amount of a compound according to any of claims 1-12.

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16. A method for the treatment or prophylaxis of cardiovascular disease in a subject, said method comprising administering to said subject a therapeutically effective amount of a compound according to any of claims 1-12.

25 17. The method according to claim 16, wherein said cardiovascular disease is selected from atherosclerosis and hypercholesterolemia.

18. A method for the treatment or prophylaxis of cholestatic liver disease in a subject comprising administering a therapeutically effective amount of a compound  
30 according to any of claims 1-12.

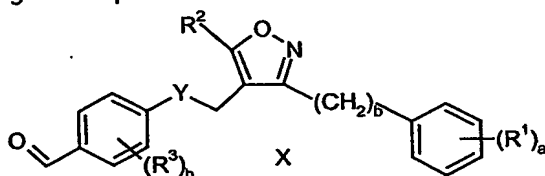
19. A method for the treatment or prophylaxis of organ fibrosis in a subject comprising administering a therapeutically effective amount of a compound according to any of claims 1-12.

5 20. A method for increasing HDL cholesterol in a subject, said method comprising administering a therapeutically effective amount of a compound according to any of claims 1-12.

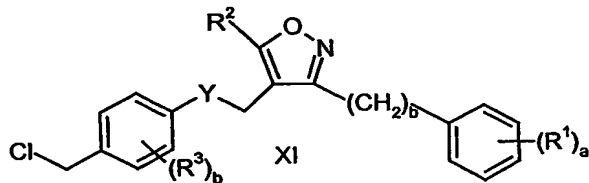
21. A method for lowering triglycerides in a subject, said method comprising  
10 administering a therapeutically effective amount of a compound according to any of claims 1-12.

22. A process for preparing a compound according to any of claims 1-12, said process comprising the steps of:

15 a) reducing a compound of formula (X):

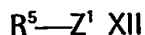


20 followed by chlorination to prepare a compound of formula (XI):



25 and

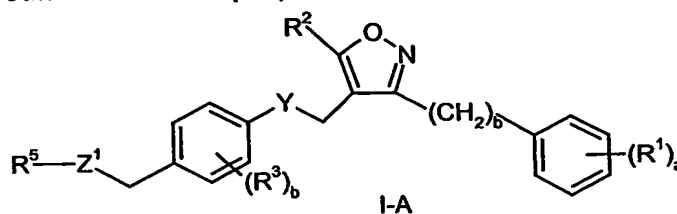
b) reacting the compound of formula (XI) with a compound of formula (XII):



wherein Z¹ is -O-, -S(O)ᵣ- or -N(R⁸)-;

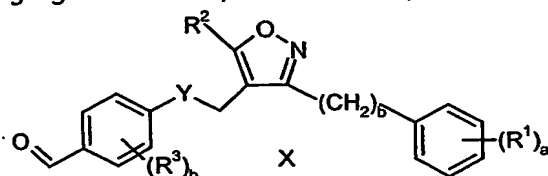
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to prepare a compound of formula (I-A):

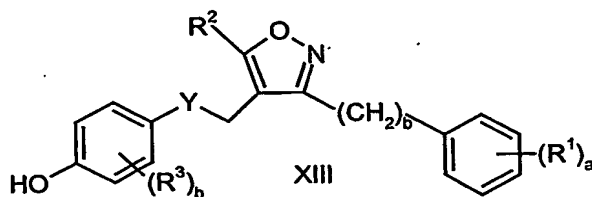


23. A process for preparing a compound according to any of claims 1-12, said process comprising the steps of:

- 10 a) rearranging the carbonyl functionality of the compound of formula (X):

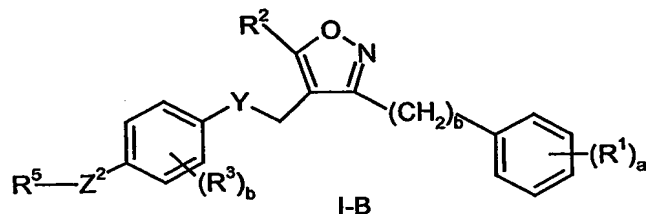


- 15 followed by hydrolysis to prepare a compound of formula (XIII):



20 and

- b) reacting the compound of formula (XIII) with a suitable electrophile to prepare a compound of formula (I-B):

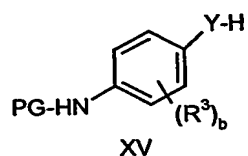


wherein Z² is -R⁴-O-.

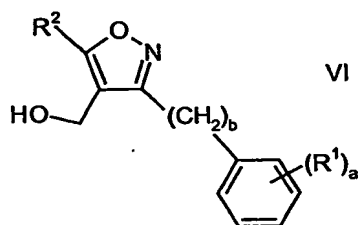
24. A process for preparing a compound according to any of claims 1-12, said process comprising the steps of:

- 30 a) reacting a protected compound of formula (XV):

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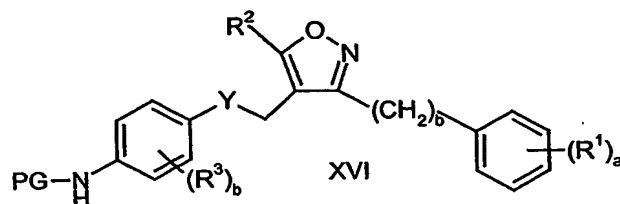


- 5        wherein PG is a protecting group;  
 with a compound of formula (VI):



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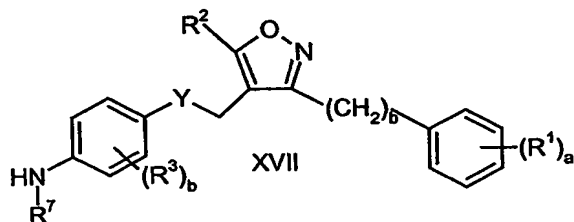
to prepare a compound of formula (XVI):



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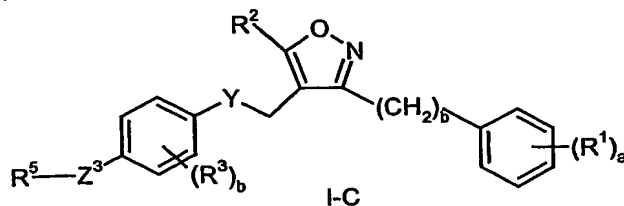
- b)        optionally alkylating the compound of formula (XVI), followed by deprotecting  
 the compound of formula (XVI) to prepare a compound of formula (XVII):

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and

- 25    c)        reacting the compound of formula (XVII) with a suitable electrophile to  
 prepare a compound of formula (I-C):



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wherein  $Z^3$  is selected from the group consisting of  $-R^4-O-$ ,  $-R^4-S(O)_r-$ ,  $-R^4-N(R^8)-$ ,  $-CON(R^8)-$ ,  $-C(O)R^4N(R^8)-$ ,  $-S(O)_rN(R^8)-$  and  $-S(O)_rR^4N(R^8)-$ .

25. A compound according to any of claims 1-12 for use in therapy.
- 5 26. A compound according to any of claims 1-12 for use in the treatment or prophylaxis of a condition mediated by FXR in a subject.
- 10 27. A compound according to any of claims 1-12 for use in the treatment or prophylaxis of cardiovascular disease in a subject.
28. A compound according to any of claims 1-12 for use in the treatment or prophylaxis of atherosclerosis or hypercholesterolemia in a subject.
- 15 29. A compound according to any of claims 1-12 for use in the treatment or prophylaxis of cholestatic liver disease in a subject.
30. A compound according to any of claims 1-12 for use in the treatment or prophylaxis of organ fibrosis in a subject.
- 20 31. A compound according to any of claims 1-12 for use in increasing HDL cholesterol in a subject.
32. A compound according to any of claims 1-12 for use in lowering triglycerides
- 25 in a subject.
33. Use of a compound according to any of claims 1-12 for the preparation of a medicament for the treatment or prophylaxis of a condition mediated by FXR in a subject.
- 30 34. Use of a compound according to any of claims 1-12 for the preparation of a medicament for the treatment or prophylaxis of cardiovascular disease in a subject.

35. Use of a compound according to any of claims 1-12 for the preparation of a medicament for the treatment or prophylaxis of atherosclerosis or hypercholesterolemia in a subject.

5 36. Use of a compound according to any of claims 1-12 for the preparation of a medicament for the treatment or prophylaxis of cholestatic liver disease in a subject.

37. Use of a compound according to any of claims 1-12 for the preparation of a medicament for the treatment or prophylaxis of organ fibrosis in a subject.

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38. Use of a compound according to any of claims 1-12 for the preparation of a medicament for increasing HDL cholesterol in a subject.

15 39. Use of a compound according to any of claims 1-12 for the preparation of a medicament for lowering triglycerides in a subject.

40. A pharmaceutical composition comprising a compound according to any of claims 1-12 for use in the treatment or prophylaxis of a condition mediated by FXR.

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